

## AMENDMENTS TO THE CLAIMS

This listing of claims will replace all prior versions of claims in the application.

### Listing of Claims:

1. (Currently amended) A purified antibody that preferentially binds a T cell antigen receptor (TCR), wherein said antibody preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of said TCR; ~~or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells.~~

2. (Original) The purified antibody of claim 1, that preferentially binds and preferentially expands an invariant T cell.

3. (Currently Amended) The purified antibody of claim 1, wherein said antibody that preferentially binds the antigen binding site of the TCR of a said T cell subpopulation.

4. (Currently amended) A composition comprising an antibody, or fragment or derivative thereof, of claim 1, said composition further comprising an antibody selected from the group consisting of an anti-V $\alpha$ 24 antibody, an anti-CD161 antibody, an anti-CD94 antibody, and an anti-V $\beta$ 11 antibody A combination of purified antibodies that preferentially binds a TCR, wherein said antibody combination preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of said TCR; ~~or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells; wherein said antibody combination is selected from the group consisting of:~~

- ~~(i) an anti-V $\alpha$ 24 antibody and an anti-CD161 antibody;~~
- ~~(ii) an anti-V $\alpha$ 24 antibody and an anti-CD94 antibody;~~

- ~~(iii) an anti-V $\beta$ 11 antibody and an anti-CD161 antibody; and~~  
~~(iv) an anti-V $\beta$ 11 antibody and an anti-CD94 antibody.~~

5. (Currently amended) A fragment or derivative of an antibody, wherein said antibody preferentially binds a CDR3-loop ~~or an  $\alpha$ - $\beta$  junction~~ of a TCR; ~~or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells.~~

6. (Withdrawn) A bifunctional antibody comprising:

(a) a first antibody or fragment thereof that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J Q<sup>+</sup> T cells; wherein said first antibody or fragment binds a first epitope; and

(b) a second antibody or fragment thereof that binds a second epitope expressed on a T cell expressing said TCR or expressed on a NK T cell, CD1d-reactive T cell, or J Q<sup>+</sup> T cell that is bound by said first antibody or fragment thereof.

7. (Currently amended) A stable hybridoma that produces an antibody, wherein said antibody preferentially binds a CDR3-loop ~~or an  $\alpha$ - $\beta$  junction~~ of a TCR; ~~or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells.~~

8. (Withdrawn) A purified T cell subpopulation, wherein said T cells are specifically bound by an antibody or a combination of antibodies, wherein said antibody or said antibody combination preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or wherein said antibody preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells.

9. (Currently amended) A method of generating the an antibody of claim 1, ~~that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup>-T cells; said method comprising:~~

- (a) coupling a cyclic peptide to a carrier;
- (b) immunizing a mammal with said coupled peptide; and
- (c) isolating an antibody of claim 1 ~~that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup>-T cells.~~

10. (Currently amended) The method of claim 9, wherein prior to step (c) said method further comprises A method of generating an antibody that ~~preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup>-T cells; said method comprising:~~

- (a) ~~immunizing the mammal a CD1 or invariant T cell deficient mammal with invariant T cells; and~~
- (b) ~~isolating an antibody that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup>-T cells.~~

11. (Currently amended) The method of claim 9 or 10, wherein said mammal is a CD1d knockout mouse, a mammal tolerized to NK T cells, a mammal tolerized to CD1d-reactive T cells, a mammal tolerized to J $\alpha$ Q<sup>+</sup> T cell, a mammal tolerized to invariant T cells, a mammal tolerized to an the invariant TCR, a mammal in which invariant T cells have been removed, a mammal lacking part of the  $\alpha$  chain of said TCR  ~~$\alpha$ -chain~~, or a mammal lacking part of the  $\beta$  chain of said TCR.

12. (Withdrawn) A method of measuring the amount of NK TCRs or the amount of NK T cells in a sample, said method comprising contacting said sample with an antibody that preferentially binds a CDR3-loop, an antigen binding site, or an  $\alpha$ - $\beta$  junction of said TCRs.

13. (Withdrawn) A method of measuring the amount of CD1d-reactive TCRs or the amount of CD1d-reactive T cells in a sample, said method comprising contacting said sample with an antibody that preferentially binds a CDR3-loop, an antigen binding site, or an  $\alpha$ - $\beta$  junction of said TCRs.

14. (Withdrawn) A method of measuring the amount of J $\alpha$ Q<sup>+</sup> TCRs or the amount of J $\alpha$ Q<sup>+</sup> T cells in a sample, said method comprising contacting said sample with an antibody or a combination of antibodies that preferentially binds a CDR3-loop, an antigen binding site, or an  $\alpha$ - $\beta$  junction of said TCRs.

15. (Withdrawn) A method of visualizing the NK TCRs or the NK T cells in a sample, said method comprising contacting said sample with an antibody that preferentially binds a CDR3-loop, an antigen binding site, or an  $\alpha$ - $\beta$  junction of said TCRs.

16. (Withdrawn) A method of visualizing the CD1d-reactive TCRs or the CD1d-reactive T cells in a sample, said method comprising contacting said sample with an antibody that preferentially binds a CDR3-loop, an antigen binding site, or an  $\alpha$ - $\beta$  junction of said TCRs.

17. (Withdrawn) A method of visualizing the J $\alpha$ Q<sup>+</sup> TCRs or the J $\alpha$ Q<sup>+</sup> T cells in a sample, said method comprising contacting said sample with an antibody or a combination of antibodies that preferentially binds a CDR3-loop, an antigen binding site, or an  $\alpha$ - $\beta$  junction of said TCRs.

18. (Withdrawn) A method of diagnosing a subject with a condition or an increased risk for a condition selected from the group consisting of autoimmune disease, viral infection, bacterial infection, parasitic infection, infection by a

eukaryotic pathogen, allergy, asthma, inflammatory condition, graft versus host disease, graft rejection, immunodeficiency disease, spontaneous abortion, pregnancy, and cancer; said method comprising the following:

(a) contacting a sample from said subject with an antibody or a combination of antibodies that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or an antibody that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and  $J\alpha Q^+$  T cells;

(b) quantitating the amount of said antibody or said antibody combination bound to said TCR or said T cells; thereby determining the amount of T cells of interest in said sample; and

(c) comparing the amount of said T cells of interest in said sample to the amount of said T cells of interest found in subjects diagnosed with said condition or subjects not diagnosed with said condition.

19. (Withdrawn) The method of claim 18, further comprising comparing the amount of another T cell type in said sample with the amount of said another T cell type found in subjects diagnosed with said condition or subjects not diagnosed with said condition.

20. (Withdrawn) A method of treating or preventing an autoimmune disease, viral infection, bacterial infection, parasitic infection, infection by a eukaryotic pathogen, allergy, asthma, inflammatory condition, graft versus host disease, graft rejection, immunodeficiency disease, spontaneous abortion, pregnancy, or cancer in a mammal, said method comprising administering to said mammal an antibody or a combination of antibodies that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and  $J\alpha Q^+$  T cells.

21. (Withdrawn) A method of inhibiting T cell pathogenesis in a mammal, said method comprising administering to said mammal an antibody or a combination of antibodies that preferentially binds a CDR3-loop, an antigen

binding site, or an  $\alpha$ - $\beta$  junction of said TCRs; or inhibits the expansion of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells; said administering sufficient to inhibit a T cell expressing said TCR, a NK T cell, a CD1d-reactive T cell, or a J $\alpha$ Q<sup>+</sup> T cell.

22. (Withdrawn) The method of claim 21, wherein said antibody is covalently linked to a toxin or a radiolabel.

23. (Currently amended) A method of increasing the size of a subpopulation of T cells, said method comprising contacting a sample comprising said T cells with an antibody or a combination of antibodies of claim 1 ~~that preferentially binds or modulates the expansion or activation of at least one T-cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, J $\alpha$ Q<sup>+</sup> T cells, and T cells expressing a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR that is preferentially bound by said antibody~~, wherein said contacting occurs under conditions that result in an increase in the number of said T cells.

24. (Original) The method of claim 23, further comprising contacting said sample with an antigen and antigen presenting cells under conditions that allow said contacting to increase the number of said T cells; wherein said antigen is not  $\alpha$ -galactosylceramide.

25. (Original) The method of claim 24, wherein said antigen is a lipid or glycosyl-phosphatidylinositol antigen from an infectious pathogen, an antigen from a cancerous cell, or a self-lipid.

26. (Original) The method of claim 23, further comprising contacting said sample with an antigen and antigen presenting cells under conditions that allow said contacting to increase the number of said T cells; wherein said antigen is  $\alpha$  - galactosylceramide.

27. (Currently amended) A method of increasing the size of a subpopulation of T cells, said method comprising:

(a) contacting a sample comprising said T cells with an antibody or a combination of antibodies of claim 1 ~~that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or an antibody that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells;~~ said contacting conducted under conditions that allow complex formation between said T cells and said antibody or said combination of antibodies;

(b) isolating said complex; and

(c) contacting said T cells in said complex or recovered from said complex with an antigen and antigen presenting cells under conditions that allow said contacting to increase the number of said T cells; wherein said antigen is not  $\alpha$ -galactosylceramide.

28. (Original) The method of claim 27, wherein said antigen is a lipid or glycosyl-phosphatidylinositol antigen from an infectious pathogen, an antigen from a cancerous cell, or a self-lipid.

29. (Currently amended) A method of increasing the size of a subpopulation of T cells, said method comprising:

(a) contacting a sample comprising said T cells with an antibody or a combination of antibodies of claim 1 ~~that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or an antibody that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells;~~ said contacting conducted under conditions that allow complex formation between said T cells and said antibody or said combination of antibodies;

(b) isolating said complex; and

(c) contacting said T cells in said complex or recovered from said complex with an antigen and antigen presenting cells under conditions that allow said contacting to increase the number of said T cells; wherein said antigen is wherein said antigen is  $\alpha$ -galactosylceramide.

30. (Original) The method of claim 27 or 29, further comprising contacting said sample or said complex with one or more cytokines.

31. (Currently amended) A method of increasing the size of a subpopulation of T cells in a mammal, said method comprising:

- (a) obtaining a sample comprising said T cells from said mammal;
- (b) contacting said T cells with an antibody or a combination of antibodies of claim 1 ~~that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, J $\alpha$ Q<sup>+</sup> T cells, and T cells expressing a CDR3 loop or an  $\alpha$ - $\beta$  junction of a TCR that is preferentially bound by said antibody or said antibody combination;~~ said contacting conducted under conditions that allow said contacting to increase the number of said T cells; and
- (c) administering said contacted T cells to said mammal.

32. (Original) The method of claim 31, further comprising purifying said T cells prior to said contacting step or after said contacting step.

33. (Currently amended) A method of increasing the size of a subpopulation of T cells in a mammal, said method comprising:

- (a) obtaining a sample comprising said T cells from said mammal;
- (b) contacting said T cells with an antibody or a combination of antibodies of claim 1 ~~that preferentially binds a CDR3 loop or an  $\alpha$ - $\beta$  junction of a TCR; or an antibody that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells;~~ said contacting conducted under conditions that allow complex formation between said T cells and said antibody or said combination of antibodies;
- (c) isolating said complex; and
- (d) contacting said T cells in said complex or recovered from said complex with an antigen and antigen presenting cells under conditions that allow said contacting to increase the number of said T cells; wherein said antigen is not  $\alpha$ -galactosylceramide; and
- (e) administering said contacted T cells to said mammal.

34. (Original) The method of claim 33, wherein said antigen is a lipid or glycosyl-phosphatidylinositol antigen from an infectious pathogen, an antigen from a cancerous cell, or a self-lipid.



35. (Currently amended) A method of increasing the size of a subpopulation of T cells in a mammal, said method comprising:

- (a) obtaining a sample comprising said T cells from said mammal;
- (b) contacting said T cells with an antibody or a combination of antibodies of claim 1 ~~that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or an antibody that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells~~; said contacting conducted under conditions that allow complex formation between said T cells and said antibody or said combination of antibodies;
- (c) isolating said complex; and
- (d) contacting said T cells in said complex or recovered from said complex with an antigen and antigen presenting cells under conditions that allow said contacting to increase the number of said T cells; wherein said antigen is  $\alpha$ -galactosylceramide; and
- (e) administering said contacted T cells to said mammal.

36. (Original) The method of claim 33 or 35, further comprising administering one or more cytokines to said mammal.

37. (Original) The method of claim 33 or 35, further comprising contacting said sample or said T cells with one or more cytokines, wherein said contacting alters the ratio of Th1/ Th2/ immune deviation response by said contacted T cells.

38. (Original) The method of claim 33 or 35, wherein said method is used in the treatment or prevention of an autoimmune disease, viral infection, bacterial infection, parasitic infection, infection by a eukaryotic pathogen, allergy, asthma, inflammatory condition, graft versus host disease, graft rejection, immunodeficiency disease, spontaneous abortion, pregnancy, or cancer in said mammal.

39. (Withdrawn) A method of purifying a subpopulation of T cells from a sample, said method comprising contacting said sample with an antibody or a combination of antibodies that preferentially binds a CDR3-loop or an  $\alpha$ - $\beta$  junction of a TCR; or an antibody that preferentially binds or modulates the expansion or activation of at least one T cell subpopulation selected from the group of NK T cells, CD1d-reactive T cells, and J $\alpha$ Q<sup>+</sup> T cells.

40. (Withdrawn) The method of claim 39, further comprising contacting said sample with an anti-V $\alpha$ 24, CD4, CD8, CD56, CD161, or V $\beta$ 11 antibody.

41. (Withdrawn) The method of claim 39, wherein said antibody is covalently linked to a fluorescent label, wherein said complex is isolated based on the fluorescence signal of said complex.

42. (Withdrawn) The method of claim 39, wherein said antibody is covalently linked to a magnetic label, wherein said complex is isolated based on the magnetism of said complex.

43. (New) The purified antibody of claim 1, wherein said antibody is 6B11 or 3A6.

44. (New) The stable hybridoma of claim 7, wherein said antibody is 6B11 or 3A6.

45. (New) The method of claim 23, wherein said antibody is 6B11 or 3A6.

46. (New) The method of claim 27, wherein said antibody is 6B11 or 3A6.

47. (New) The method of claim 29, wherein said antibody is 6B11 or 3A6.

48. (New) The method of claim 31, wherein said antibody is 6B11 or 3A6.

49. (New) The method of claim 33, wherein said antibody is 6B11 or 3A6.

50. (New) The method of claim 35, wherein said antibody is 6B11 or 3A6.

51. (New) The method of claim 9, wherein said mammal is a CD1 or invariant T cell deficient mammal.

52. (New) The purified antibody of claim 1, wherein said antibody preferentially binds at least one T cell population selected from the group consisting of NK T cells, CD1d reactive T cells, and JαQ<sup>+</sup> T cells.

53. (New) The purified antibody of claim 1, wherein said antibody preferentially modulates the expansion or activation of at least one T cell population selected from the group consisting of NK T cells, CD1d reactive T cells, and JαQ<sup>+</sup> T cells.

54. (New) The purified antibody of claim 1, wherein said antibody preferentially binds an invariant T cell.

55. (New) The purified antibody of claim 1, wherein said antibody is a monoclonal antibody.

56. (New) The purified antibody of claim 1, wherein said antibody is humanized.

57. (New) The fragment or derivative of an antibody of claim 5, wherein said fragment or derivative preferentially binds at least one T cell population selected from the group consisting of NK T cells, CD1d reactive T cells, and JαQ<sup>+</sup> T cells.

58. (New) The fragment or derivative of an antibody of claim 5, wherein said fragment or derivative preferentially modulates the expansion or activation of at least one T cell population selected from the group consisting of NK T cells, CD1d reactive T cells, and JαQ<sup>+</sup> T cells.

59. (New) The fragment or derivative of an antibody of claim 5, wherein said

fragment or derivative preferentially binds an invariant T cell.

60. (New) The fragment or derivative of an antibody of claim 5, wherein said antibody is a monoclonal antibody.

61. (New) The fragment or derivative of an antibody of claim 5, wherein said fragment or derivative is humanized.

62. (New) The fragment or derivative of an antibody of claim 5, wherein said fragment is a ScFv, Fab, or F(ab')<sub>2</sub> fragment.

63. (New) The fragment or derivative of an antibody of claim 5, wherein said derivative is an antibody linked to a toxin, a therapeutically active compound, an enzyme, a cytokine, a radiolabel, a fluorescent label, a magnetic label, or an affinity tag.

64. (New) The stable hybridoma of claim 7, wherein said antibody preferentially binds at least one T cell population selected from the group consisting of NK T cells, CD1d reactive T cells, and JαQ<sup>+</sup> T cells.

65. (New) The stable hybridoma of claim 7, wherein said antibody preferentially modulates the expansion or activation of at least one T cell population selected from the group consisting of NK T cells, CD1d reactive T cells, and JαQ<sup>+</sup> T cells.

66. (New) The stable hybridoma of claim 7, wherein said antibody preferentially

binds an invariant T cell.

67. (New) The stable hybridoma of claim 66, wherein said antibody preferentially expands said invariant T cell.

68. (New) The stable hybridoma of claim 7, wherein said antibody is a monoclonal antibody.